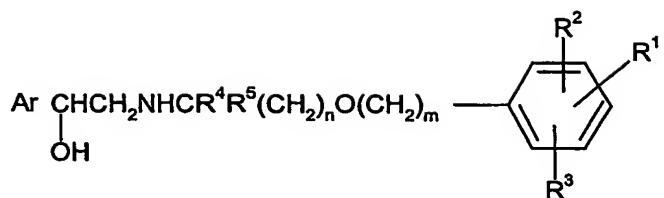


## CLAIMS

## 1. A compound of formula (I)

5



(I)

or a salt, solvate, or physiologically functional derivative thereof, wherein:

10 n is an integer of from 2 to 8;

m is an integer of from 3 to 11, with the proviso that the sum of n + m is from 5 to 19;

R<sup>1</sup> is hydrogen or -XSO<sub>2</sub>NR<sup>6</sup>R<sup>7</sup>;

15 wherein X is -(CH<sub>2</sub>)<sub>p</sub>- or C<sub>2-6</sub> alkenylene;

p is an integer from 0 to 6;

20 R<sup>6</sup> and R<sup>7</sup> are independently selected from hydrogen, C<sub>1-6</sub>alkyl, C<sub>3-7</sub>cycloalkyl, CONR<sup>8</sup>R<sup>9</sup>, phenyl and phenyl(C<sub>1-4</sub>alkyl)-,

or R<sup>6</sup> and R<sup>7</sup>, together with the nitrogen atom to which they are bonded, form a 5-, 6- or 7-membered nitrogen – containing ring;

25 and R<sup>6</sup> and R<sup>7</sup> are each independently optionally substituted by 1 or 2 groups independently selected from halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkoxy, hydroxy-substituted C<sub>1-6</sub>alkoxy, C<sub>1-6</sub>haloalkyl, CO<sub>2</sub>R<sup>8</sup>, SO<sub>2</sub>R<sup>8</sup>R<sup>9</sup>, -CONR<sup>8</sup>R<sup>9</sup>, -NR<sup>8</sup>C(O)R<sup>9</sup> or a 5-, 6- or 7- membered heterocyclic ring;

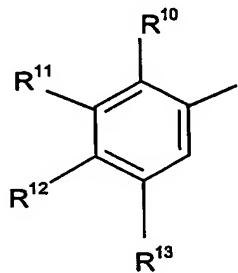
$R^8$  and  $R^9$  are independently selected from hydrogen,  $C_{1-6}$ alkyl,  $C_{3-7}$ cycloalkyl, phenyl and phenyl( $C_{1-6}$ alkyl)-;

$R^2$  and  $R^3$  are independently selected from hydrogen,  $C_{1-6}$ alkyl,  $C_{1-6}$ alkoxy, halo, phenyl and  $C_{1-6}$ haloalkyl;

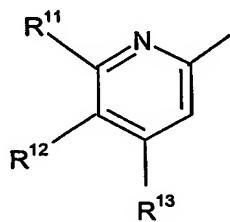
$R^4$  and  $R^5$  are independently selected from hydrogen and  $C_{1-4}$  alkyl with the proviso that the total number of carbon atoms in  $R^4$  and  $R^5$  is not more than 4,

10 and

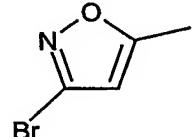
Ar is a group selected from



(a)

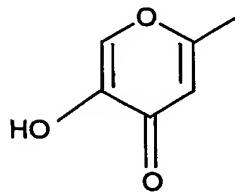


(b)



(c)

and



(d)

15

wherein  $R^{11}$  represents hydrogen, halogen,  $-(CH_2)_qOR^{14}$ ,  $-NR^{14}C(O)R^{15}$ ,  $-NR^{14}SO_2R^{15}$ ,  $-SO_2NR^{14}R^{15}$ ,  $-NR^{14}R^{15}$ ,  $-OC(O)R^{16}$  or  $OC(O)NR^{14}R^{15}$ ,

and  $R^{10}$  represents hydrogen, halogen or  $C_{1-4}$  alkyl;

or  $R^{11}$  represents  $-NHR^{17}$  and  $R^{10}$  and  $-NHR^{17}$  together form a 5- or 6- membered heterocyclic ring;

5

$R^{12}$  represents hydrogen, halogen,  $-OR^{14}$  or  $-NR^{14}R^{15}$ ;  $-OC(O)R^{16}$  or  $-OC(O)NR^{14}R^{15}$ ;

$R^{13}$  represents hydrogen, halogen,  $haloC_{1-4}$  alkyl,  $-OR^{14}$  or  $-NR^{14}R^{15}$ ;

10  $R^{14}$  and  $R^{15}$  each independently represents hydrogen or  $C_{1-4}$  alkyl, or in the groups  $-NR^{14}R^{15}$ ,  $-SO_2NR^{14}R^{15}$  and  $-OC(O)NR^{14}R^{15}$ ,  $R^{14}$  and  $R^{15}$  independently represent hydrogen or  $C_{1-4}$  alkyl or together with the nitrogen atom to which they are attached form a 5-, 6- or 7- membered nitrogen-containing ring,

15  $R^{16}$  represents an aryl (eg phenyl or naphthyl) group which may be unsubstituted or substituted by one or more substituents selected from halogen,  $C_{1-4}$  alkyl, hydroxy,  $C_{1-4}$  alkoxy or halo  $C_{1-4}$  alkyl; and

q is zero or an integer from 1 to 4;

20

provided that when  $R^1$  is hydrogen

Ar is not a group (a) wherein;

$R^{11}$  is  $-(CH_2)_qOR^{14}$ , q is zero or 1 and  $R^{12}$  is  $OR^{14}$ ,

or  $R^{11}$  is  $-(CH_2)_qOR^{14}$ , q is zero and  $R^{13}$  is  $OR^{14}$ ,

25 or  $R^{11}$  is  $-NR^{14}SO_2R^{15}$  or  $NR^{14}COR^{15}$  and  $R^{12}$  is  $OR^{14}$ ,

or  $R^{11}$  and  $R^{13}$  both represent halogen and  $R^{12}$  is  $NR^{14}R^{15}$ ;

Ar is not a group (b) wherein  $R^{11}$  is  $-(CH_2)_qOR^{14}$  and  $R^{12}$  is  $OR^{14}$ ;

Ar is not a group (c),

and when  $R^1$  is  $XSO_2NR^6R^7$ , Ar is not a group (a) wherein

30  $R^{11}$  is  $(CH_2)_qOR^{14}$  or  $NR^{14}COR^{15}$ , and  $R^{12}$  is  $OR^{14}$ .

2. A compound of formula (I) according to claim 1 wherein, in the group Ar,  $R^{11}$  represents halogen,  $-(CH_2)_qOR^{14}$ ,  $-NR^{14}C(O)R^{15}$ ,  $-NR^{14}SO_2R^{15}$ ,  $-SO_2NR^{14}R^{15}$ ,  $-NR^{14}R^{15}$ ,  $-OC(O)R^{16}$  or  $OC(O)NR^{14}R^{15}$ ,

35

and R<sup>10</sup> represents hydrogen,

or R<sup>11</sup> represents -NHR<sup>17</sup> and R<sup>10</sup> and -NHR<sup>17</sup> together form a 5- or 6- membered heterocyclic ring;

5

and

R<sup>13</sup> represents hydrogen, halogen, haloC<sub>1-4</sub> alkyl, -OR<sup>14</sup>, or -NR<sup>14</sup>R<sup>15</sup>,

10 and all other substituents are as defined in claim 1.

3. A compound of formula (I) according to claim 1 or claim 2 wherein the group R<sup>1</sup> is attached to the meta-position relative to the -O-(CH<sub>2</sub>)<sub>m</sub> link.

15 4. A compound of formula (I) according to any of claims 1 to 3 wherein R<sup>1</sup> represents SO<sub>2</sub>NR<sup>6</sup>R<sup>7</sup> wherein R<sup>6</sup> and R<sup>7</sup> are independently selected from hydrogen and C<sub>1-6</sub>alkyl.

5. A compound of formula (I) according to any of claims 1 to 4 wherein R<sup>4</sup> and R<sup>5</sup> are independently selected from hydrogen and methyl.

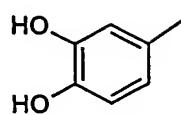
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6. A compound of formula (I) according to any of claims 1 to 5 wherein R<sup>2</sup> and R<sup>3</sup> each represent hydrogen.

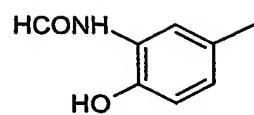
25 7. A compound of formula (I) according to any of claims 1 to 6 wherein n is 5 or 6 and m is 3 or 4 such that m + n is 8, 9 or 10.

8. A compound of formula (I) according to any of claims 1 to 7 wherein Ar represents a group selected from:

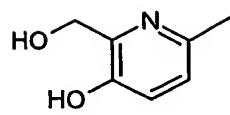
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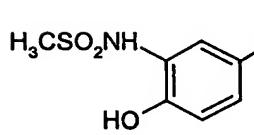
(i)



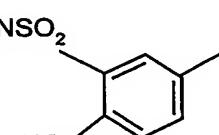
(ii)



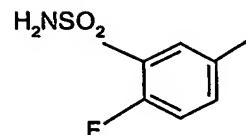
(iii)



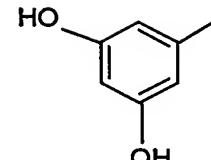
(iv)



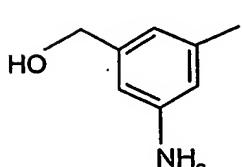
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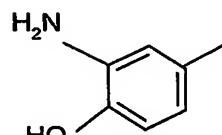
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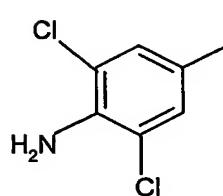
(vii)



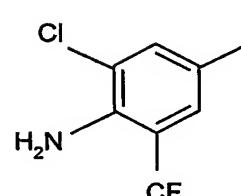
(viii)



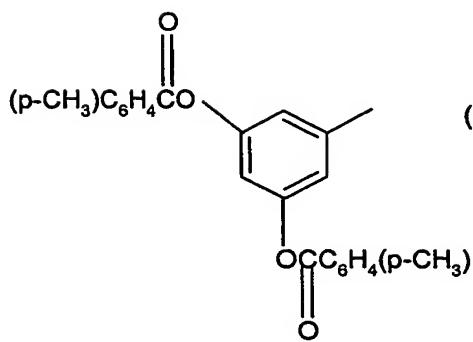
(ix)



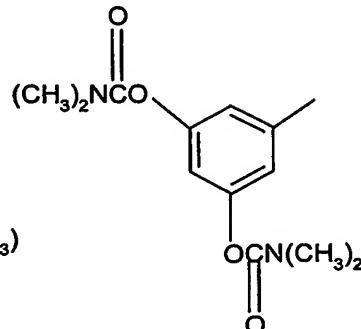
(x)



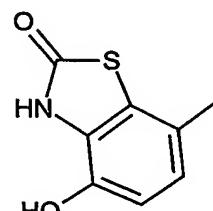
(xi)



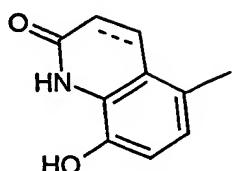
(xii)



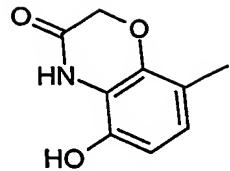
(xiii)



(xiv)



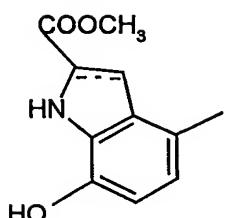
(xv)



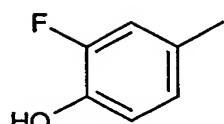
(xvi)



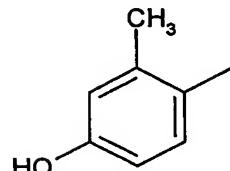
(xvii)



(xviii)



(xix)



(xx)

5 9. A compound of formula (I) according to any of claims 1 to 8 wherein R<sup>1</sup> is  
 10 hydrogen and Ar is selected from a group of structure (ii), (v), (vi), (viii), (ix), (xi), (xii), (xiii),  
 (xiv), (xv), (xvi), (xvii) and (xviii).

10. A compound of formula (I) according to any of claims 1 to 8 wherein R<sup>1</sup> is  
 15 XSO<sub>2</sub>NR<sup>6</sup>R<sup>7</sup> and Ar is selected from a group of structure (iii), (iv), (xiv), (xv), (xvi) and (xix)

11. A compound of formula (I) selected from:  
 8-Hydroxy-5-((1R)-1-hydroxy-2-((6-(4-phenylbutoxy)hexyl)amino)ethyl)quinolin-2(1H)-one;  
 3-{4-[(6-[(2R)-2-Hydroxy-2-(8-hydroxy-2-oxo-1,2-dihydroquinolin-5-  
 15 yl)ethyl]amino)hexyl]oxy}butyl}benzenesulfonamide;  
 5-Hydroxy-8-(1-hydroxy-2-((6-(4-phenylbutoxy)hexyl)amino)ethyl)-2H-1,4-benzoxazin-  
 3(4H)-one;  
 3-{4-[(6-{[2-hydroxy-2-(5-hydroxy-3-oxo-3,4-dihydro-2H-1,4-benzoxazin-8-  
 yl)ethyl]amino}hexyl)oxy}butyl}benzenesulfonamide;

4-Hydroxy-7-((1*R*)-1-hydroxy-2-{[6-(4-phenylbutoxy)hexyl]amino}ethyl)-1,3-benzothiazol-2(3*H*)-one;

4-Hydroxy-7-(1-hydroxy-2-{[6-(4-phenylbutoxy)hexyl]amino}ethyl)-1,3-benzothiazol-2(3*H*)-one;

5 3-{4-[(6-((2*R*)-2-(3-Fluoro-4-hydroxyphenyl)-2-hydroxyethyl]amino)hexyl]oxy]butyl}benzenesulfonamide;  
3-(4-[(6-((2*R*)-2-(3-Fluoro-4-hydroxyphenyl)-2-hydroxyethyl]amino)hexyl]oxy)butyl)benzenesulfonamide;  
3-[4-((6-((2*R*)-2-Hydroxy-2-{4-hydroxy-3-

10 [(methylsulfonyl)amino]phenyl]ethyl)amino]hexyl]oxy)butyl}benzenesulfonamide;  
3-{3-[(7-((2*R*)-2-(3-Fluoro-4-hydroxyphenyl)-2-hydroxyethyl]amino)heptyl]oxy}propyl}benzenesulfonamide;  
3-(3-[(7-((2*R*)-2-Hydroxy-2-{5-hydroxy-6-(hydroxymethyl)pyridin-2-yl}ethyl)amino)heptyl]oxy)propyl}benzenesulfonamide;

15 3-[3-((7-((2*R*)-2-Hydroxy-2-{4-hydroxy-3-[(methylsulfonyl)amino]phenyl}ethyl)amino)heptyl]oxy)propyl}benzenesulfonamide;  
3-{3-[(7-((2*R*)-2-Hydroxy-2-(8-hydroxy-2-oxo-1,2-dihydroquinolin-5-yl)ethyl)amino)heptyl]oxy}propyl}benzenesulfonamide;  
3-(3-[(7-((2*R*)-2-[3-(Formylamino)-4-hydroxyphenyl]-2-hydroxyethyl)amino)heptyl]oxy)propyl}benzenesulfonamide;

20

or a salt, solvate or physiologically functional derivative thereof.

12. A method for the prophylaxis or treatment of a clinical condition in a mammal, such as a human, for which a selective  $\beta_2$ -adrenoreceptor agonist is indicated, which comprises administration of a therapeutically effective amount of a compound of formula (I) according to any of claims 1 to 11, or a pharmaceutically acceptable salt, solvate, or physiologically functional derivative thereof.

30 13. A compound of formula (I), according to any of claims 1 to 11, or a pharmaceutically acceptable salt, solvate, or physiologically functional derivative thereof for use in medical therapy.

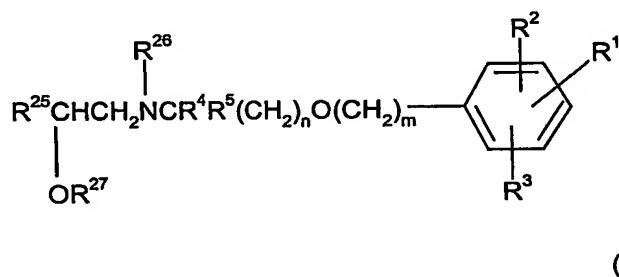
14. A pharmaceutical formulation comprising a compound of formula (I), according to any of claims 1 to 11, or a pharmaceutically acceptable salt, solvate, or physiologically

functional derivative thereof, and a pharmaceutically acceptable carrier or excipient, and optionally one or more other therapeutic ingredients.

15. The use of a compound of formula (I), according to any of claims 1 to 11, or a  
5 pharmaceutically acceptable salt, solvate, or physiologically functional derivative thereof  
in the manufacture of a medicament for the prophylaxis or treatment of a clinical condition  
for which a selective  $\beta_2$ -adrenoreceptor agonist is indicated.

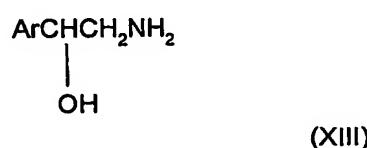
16. A process for the preparation of a compound of formula (I), according to any of  
10 claims 1 to 11, or a salt, solvate, or physiologically functional derivative thereof, which  
comprises:

(a) deprotection of a protected intermediate of formula (II):

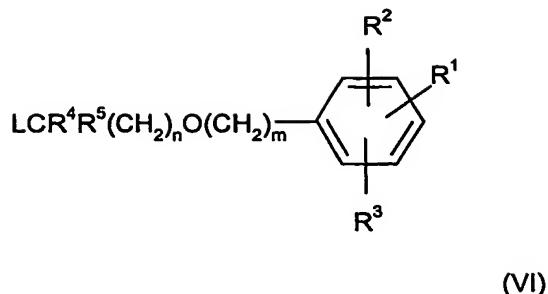


15 or a salt or solvate thereof, wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, m and n are as defined for the  
compounds of formula (I) R<sup>25</sup> represents an optionally protected form of Ar, and R<sup>26</sup> and  
R<sup>27</sup> each independently represent either hydrogen or a protecting group, provided that the  
compound of formula (II) contains at least one protecting group;

20 (b) reacting a compound of formula (XIII):



Wherein Ar is as defined above with a compound of formula (VI):



Wherein L is a leaving group such as halo (typically chloro, bromo or iodo) or a sulphonate (typically methanesulphonate) and R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, n and m are as defined

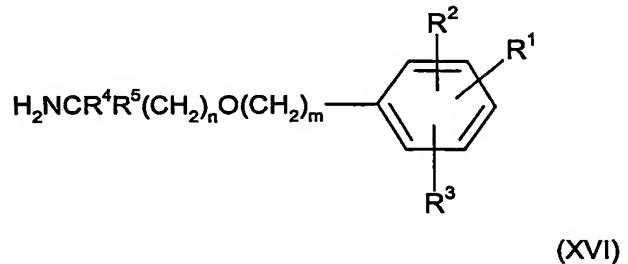
5 for compounds of formula (I).

(c) reacting a compound of formula (XV):



10

wherein L is a leaving group as hereinbefore defined, with an amine of formula (XVI):

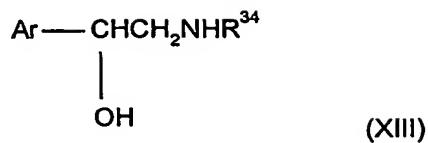


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wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, R<sup>5</sup>, n and m are as defined for formula (I); and

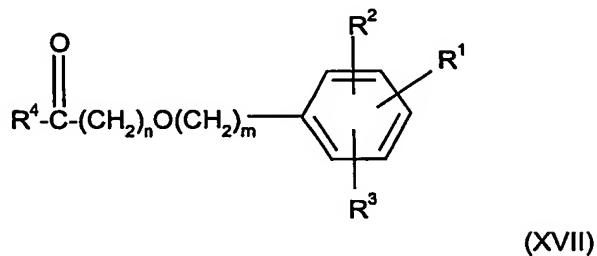
(d) (i) reacting a compound of formula (XIII):

20



Wherein Ar is as hereinbefore defined and R<sup>34</sup> is a chiral auxiliary group,

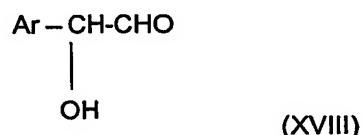
5 with a compound of formula (XVII):



wherein R<sup>1</sup>, R<sup>2</sup>, R<sup>3</sup>, R<sup>4</sup>, n and m are as hereinbefore defined;  
followed where necessary by removal of said chiral auxiliary group R<sup>34</sup>;

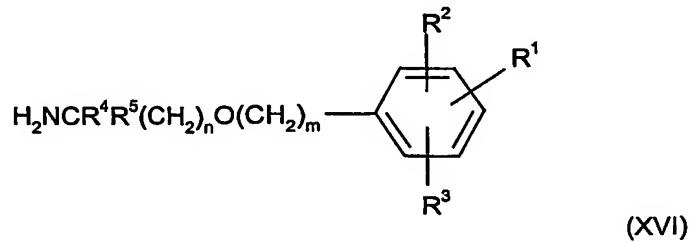
10

or (ii) reacting a compound of formula (XVIII):



wherein Ar is as hereinbefore defined; with an amine of formula (XVI):

15



as hereinbefore defined,

5 under conditions suitable to effect reductive amination,

followed by the following steps in any order:

(i) optional removal of any protecting groups;

10 (ii) optional separation of an enantiomer from a mixture of enantiomers;

(iii) optional conversion of the product to a corresponding salt, solvate,

(iv) optional conversion of a group R<sup>1</sup>, R<sup>2</sup> and/or R<sup>3</sup> to another group R<sup>1</sup>, R<sup>2</sup> and/or R<sup>3</sup>,

or physiologically functional derivative thereof.

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